

Evaluation of whole genome sequencing for routine subtyping of *Salmonella* Typhimurium for surveillance and outbreak investigation

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CONTEXT

Subtyping = characterisation of bacterial isolates beyond the species and subspecies level

- Why?
- Up-to-date diagnosis and surveillance of infectious diseases
 - Outbreak: identify a link between the origin of the infection and the human isolate

Case study: *Salmonella enterica* subsp. *enterica* serovar Typhimurium (*S.* Typhimurium) and serovar 1,4,[5],12:i:- (*S.* 1,4,[5],12:i:-)
→ important foodborne pathogen

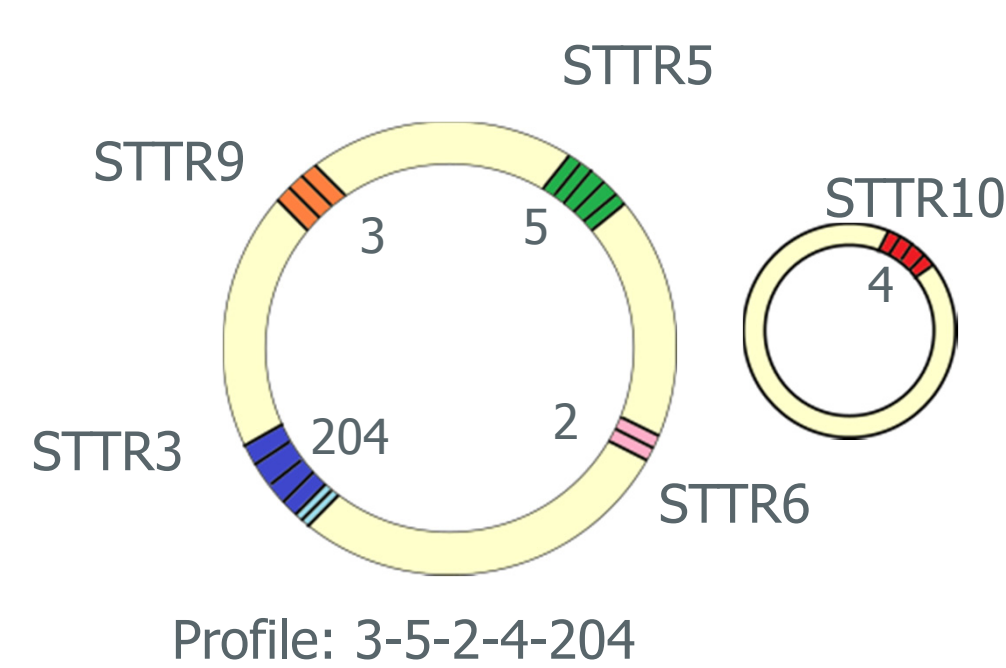
CURRENT SITUATION @ BELGIAN NRC FOR SUBTYPING

Phage typing



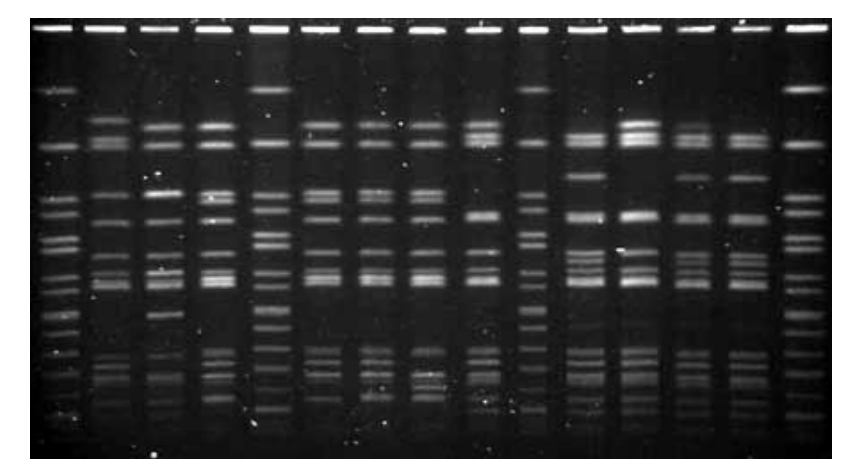
- ⊕ Very discriminative
Cheap
- ⊖ High level of expertise
NT and RDNC isolates

MLVA



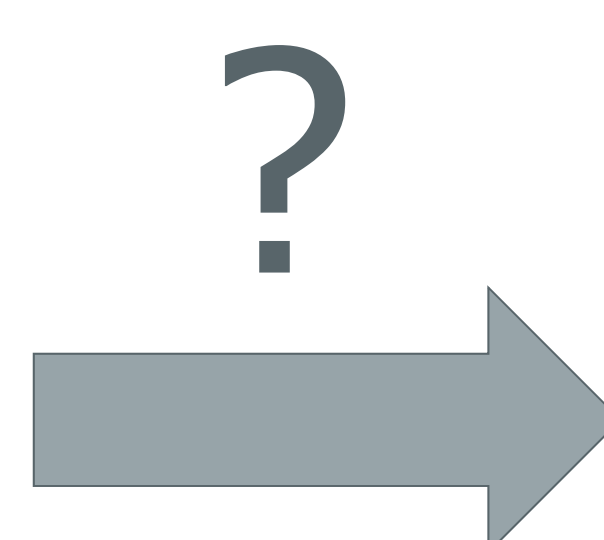
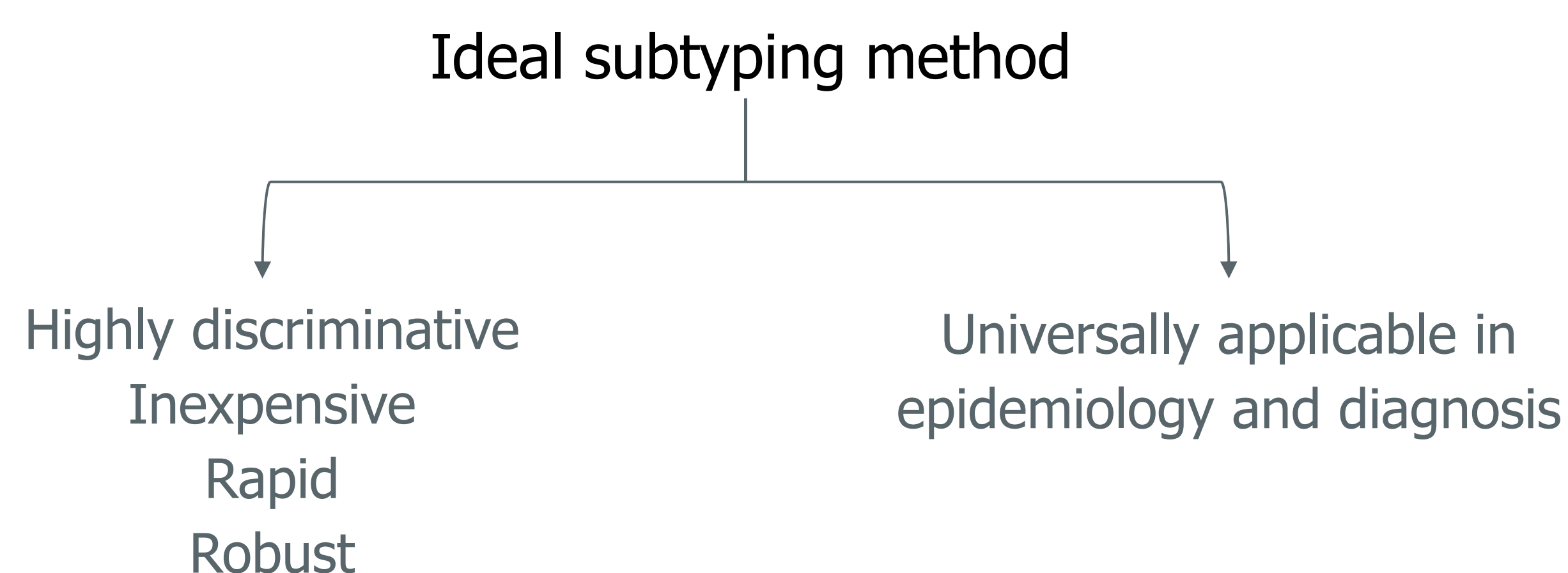
- ⊕ Profiles easily compared between labs
Rapid
- ⊖ Too many profiles for surveillance
STTR5, STTR6, STTR10 not stable → outbreak?
(Wuyts *et al.*, 2013; Dimovski *et al.*, 2014)

PFGE



- ⊕ Medium discriminative
- ⊖ Slow
Limited reproducibility

IDEAL SITUATION @ BELGIAN NRC FOR SUBTYPING



Whole genome
sequencing

ONGOING: EVALUATION OF WHOLE GENOME SEQUENCING FOR ROUTINE SUBTYPING

Human *S.* Typhimurium and *S.* 1,4,[5],12:i:- isolates

Serovar	MLVA	Phage type					RDNC
		DT104L	DT120	DT138	DT193		
1,4,[5],12:i:-	3-12-10-NA-211	-	4	-	8	-	-
	3-12-11-NA-211	-	1	10	-	5	-
	3-13-11-NA-211	-	-	5	3	2	-
	3-14-11-NA-211	-	-	-	2	-	-
Typhimurium	3-12-10-NA-211	-	7	1	3	-	-
	3-12-11-NA-211	-	11	-	3	1	-
	3-13-11-NA-211	-	1	-	-	-	-
	3-14-11-NA-211	-	-	-	-	1	-
	3-14-18-14-311	1	-	-	-	-	-
	3-16-16-13-311	2	-	-	-	-	-

Outbreak-related → 7 isolates
3 (or 1) isolates → 7×3+1 = 22 isolates
29 isolates

Data generation: next generation sequencing

- Illumina in outsourcing
- ↳ Library type?
- ↳ Coverage?

Data analysis and interpretation

- Workflow: SNP- or allele-based?
- Definition of distinct subtype?
- Genotype – phenotype?
- ↳ Antibiotic resistance
- Link with historical data?
- ↳ Phage type – MLVA

Universal subtyping method

- Other pathogens
- ↳ e.g. *S.* Enteritidis

Applicable in a
routine setting?